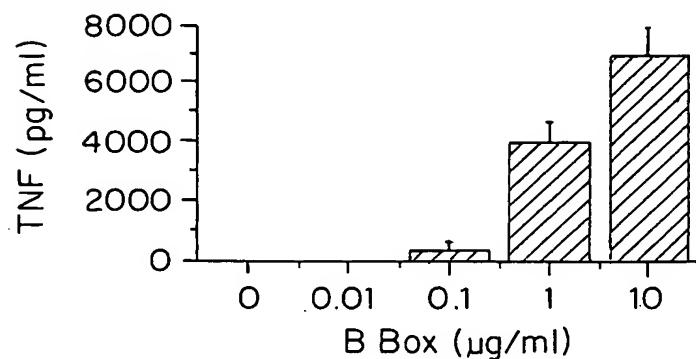
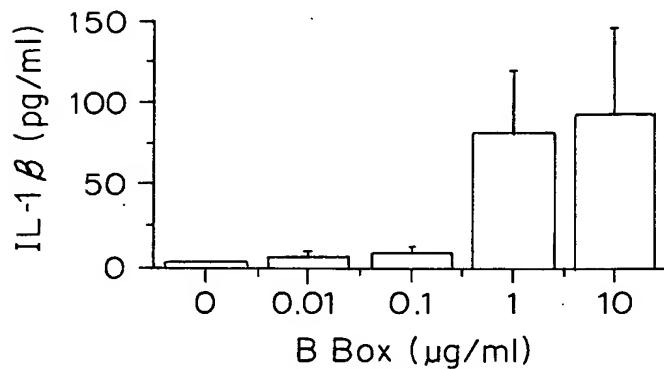


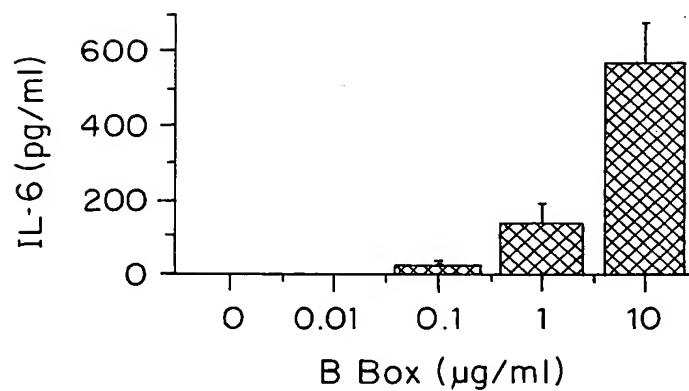
**FIG. I**



**FIG. 2A**

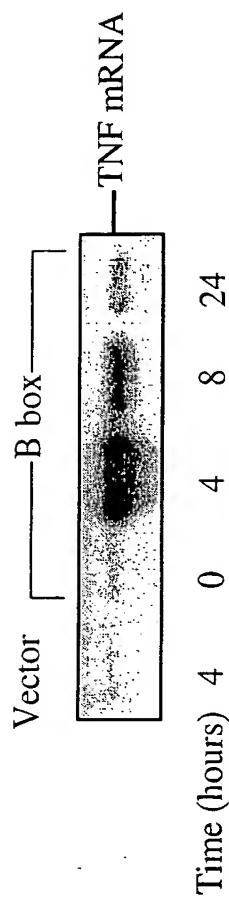


**FIG. 2B**



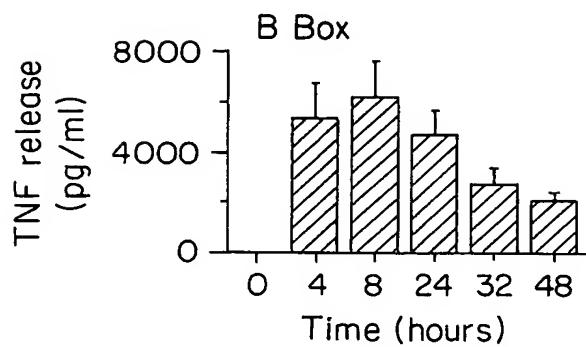
**FIG. 2C**

Docket/App No.: 3258.1008-001  
Title: HMGB1 Combination Therapies  
Inventors: Walter Newman

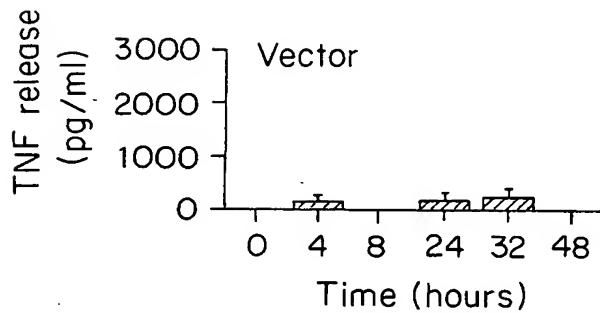


**BEST AVAILABLE COPY**

FIG. 2D



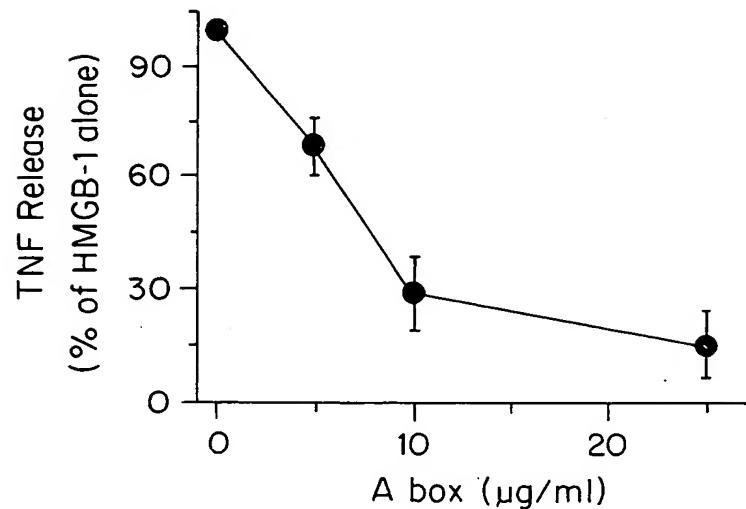
**FIG. 2E**



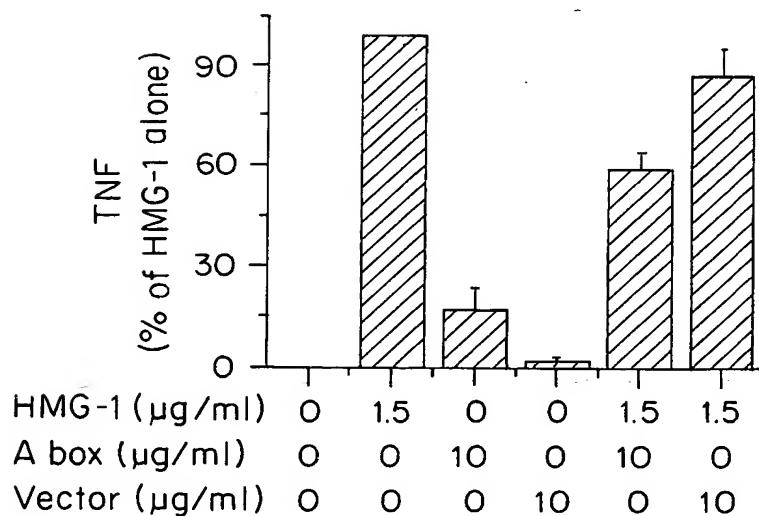
**FIG. 2F**

B box mutants	TNF release (pg/ml)
B box: 74 amino acids	5675±575
1-20	2100±756
16-35	100±10
30-49	120±75
45-64	100±36
60-74	100±20

**FIG. 3**



**FIG. 4A**



**FIG. 4B**

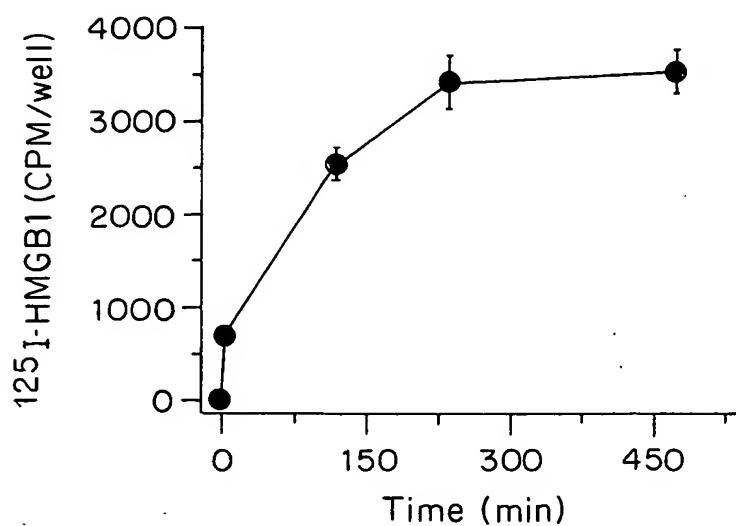


FIG. 5A

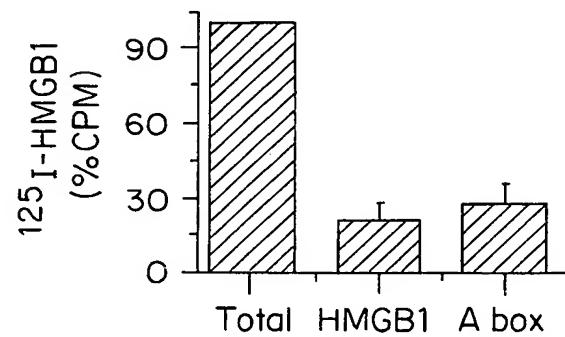


FIG. 5B

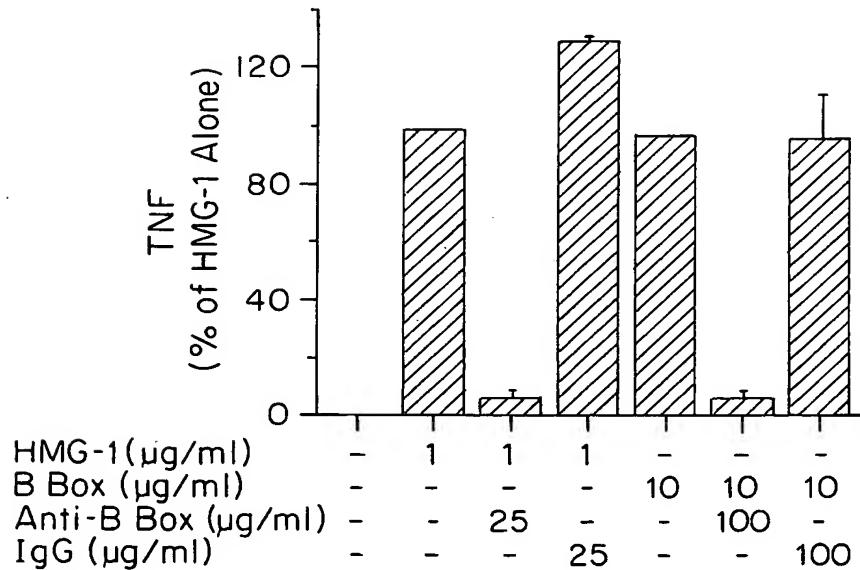


FIG. 6

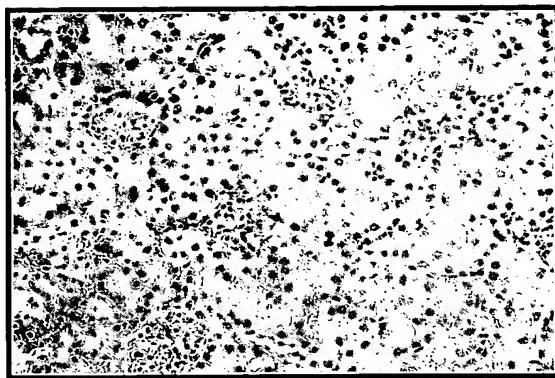


FIG. 7A

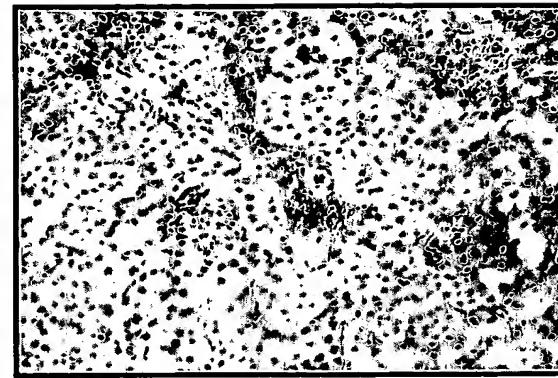


FIG. 7B

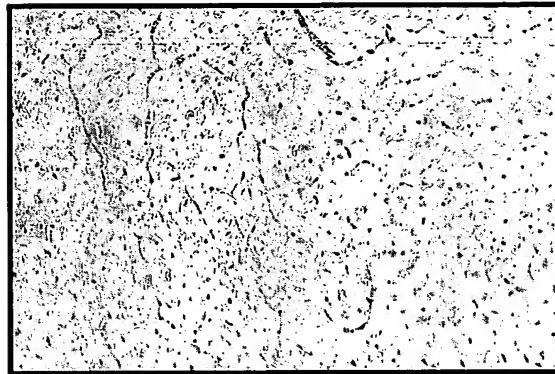


FIG. 7C



FIG. 7D



FIG. 7E

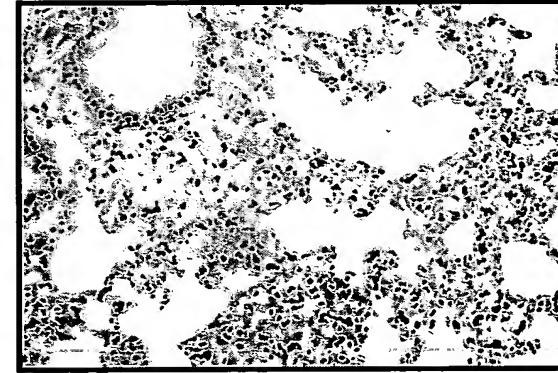
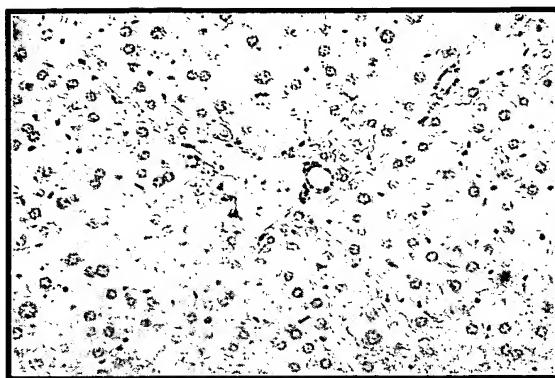
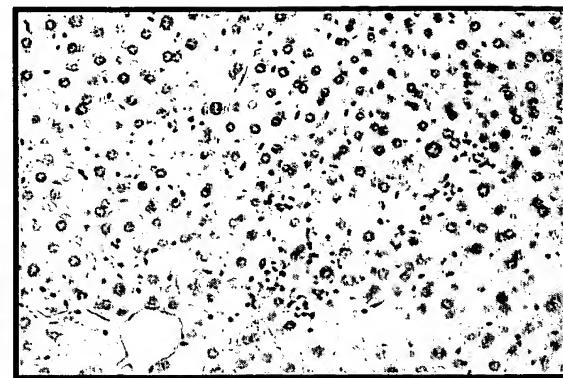


FIG. 7F

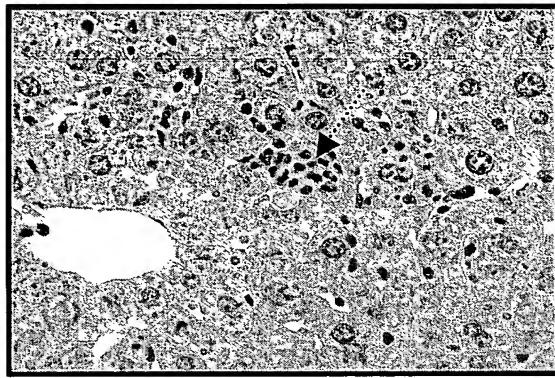
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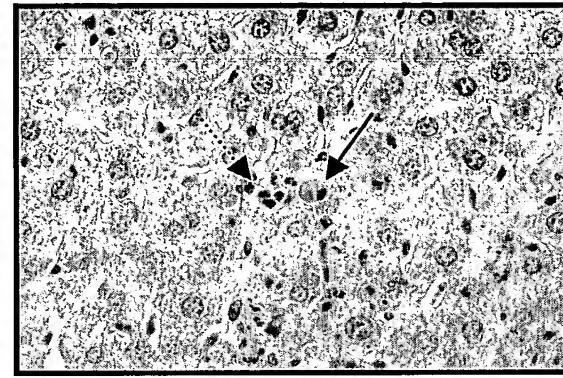
**FIG. 7G**



**FIG. 7H**

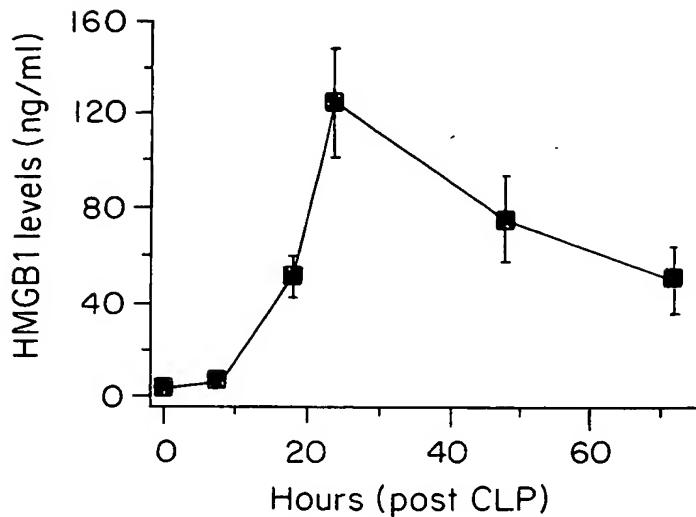


**FIG. 7I**

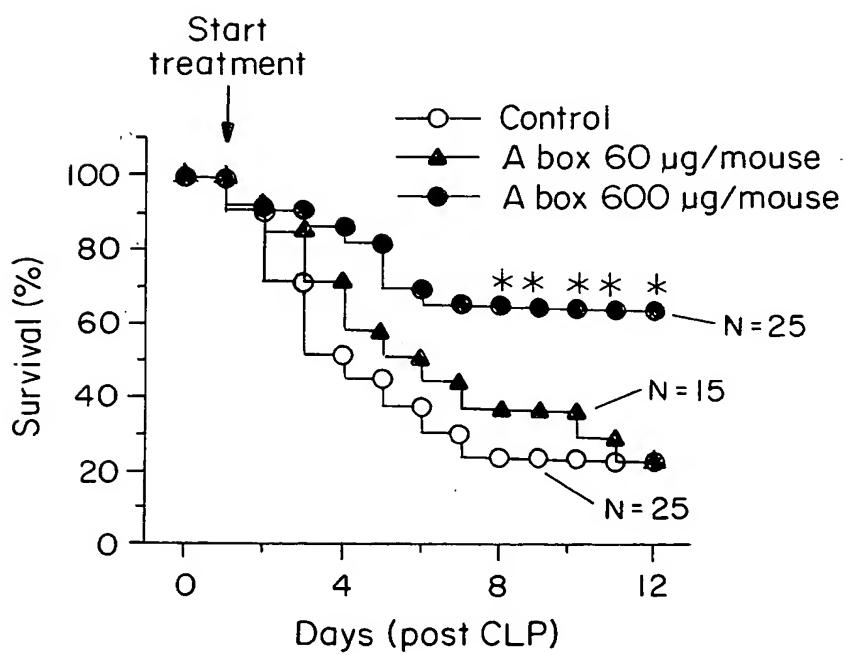


**FIG. 7J**

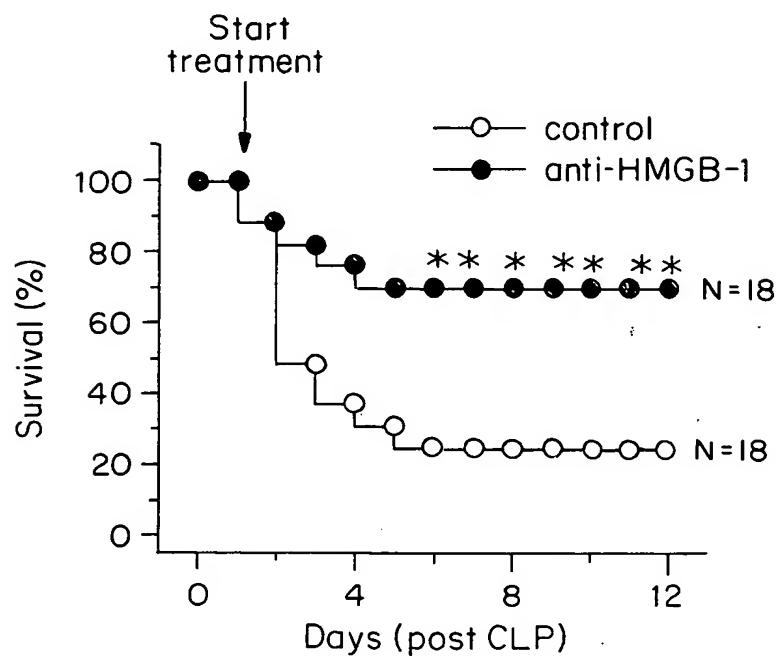
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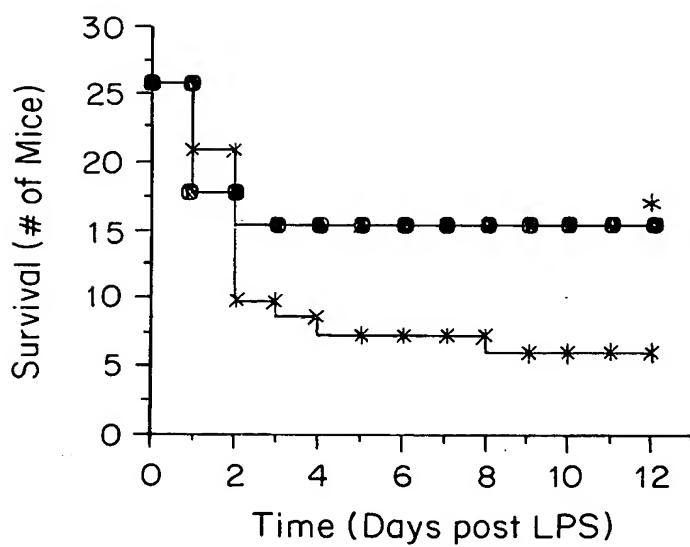
**FIG. 8**



**FIG. 9**



**FIG. 10A**



**FIG. 10B**

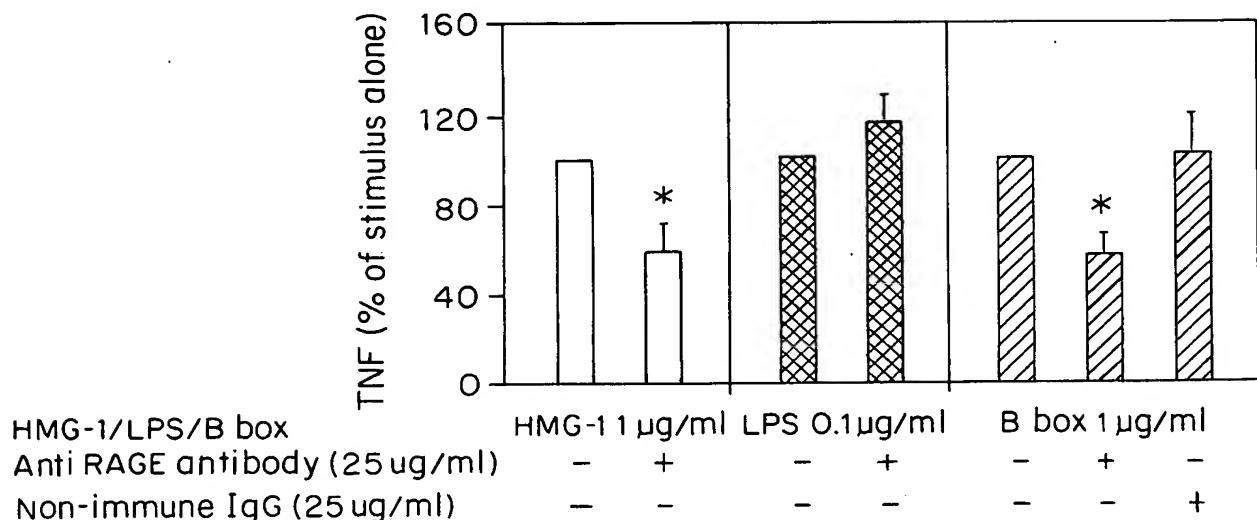


FIG. II A

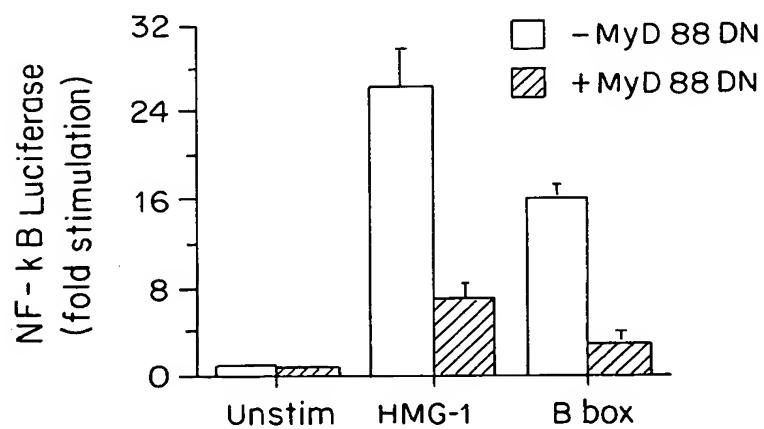


FIG. II B

FIG. 12A

SEQ ID NO:1 - Human HMG1 amino acid sequence

1 mgkgdpkkpr gkmssyaffv qtcreehkkk hpdasvnse fskkcserwk trnsakekgkf  
61 edmakadkar yeremktyip pkgetkkkfk dnapkrpps afflfcseyr pkikgehpgl  
121 sigdvakklg emwnntaadd kqpyekkaak lkekyekdia ayrakgkpda akkgvvkaek  
181 skkkkeeed eeedeedeee edeededeee dddde

FIG. 12B

SEQ ID NO:2 - Mouse and Rat HMG1 amino acid sequence

1 mgkgdpkkpr gkmssyaffv qtcreehkkk hpdasvnse fskkcserwk trnsakekgkf  
61 edmakadkar yeremktyip pkgetkkkfk dnapkrpps afflfcseyr pkikgehpgl  
121 sigdvakklg emwnntaadd kqpyekkaak lkekyekdia ayrakgkpda akkgvvkaek  
181 skkkkeeedd eeedeedeee eeeeededeee dddde

FIG. 12C

SEQ ID NO:3 - HUMAN HMG2 amino acid sequence

1 mgkgdpnkpr gkmssyaffv qtcreehkkk hpdsvnfae fskkcserwk trnsakekskf  
61 edmakadkar ydremknyvp pkgdkggkkk dnapkrpps afflfcsehr pkiksehpgl  
121 sigdtakklg emwseqsakd kqpyeqkaak lkekyekdia ayrakgksea gkkpgprtgc  
181 skkknepede eeeeeeeded eeeeededee

FIG. 12D

SEQ ID NO:4 - Human, mouse and rat HMG1 A box protein sequence

1 pdasvnsef skkcserwkt msakekgkfe dmakadkary eremktyipp kget

FIG. 12E

SEQ ID NO:5 - Human, mouse and rat HMG1 B box protein sequence

1 napkrppsaf flfcseyrpk ikgehpglsi gdvakklgem wnntaaddkq pyekkaaklk  
61 ekyekdiaa

FIG. 12F

SEQ ID NO:6 - forward PCR primer for human HMG1

gatgggcaaaggagatcctaag.

FIG. 12G

SEQ ID NO:7 - reverse PCR primer for human HMG1

gcggccgcttattcatcatcatcatcttc

FIG. 12H

SEQ ID NO:8 - forward PCR primer for -C mutant of human HMG1

gatgggcaaaggagatcctaag

FIG. 12I

SEQ ID NO:9 - reverse PCR primer for -C mutant of human HMG1  
gcggccgctcacttgcttttcagccttgac

FIG. 12J

SEQ ID NO:10 - forward PCR primer for A+B boxes mutant of human HMG1  
gagcataagaagaagcacca

FIG. 12K

SEQ ID NO:11 - reverse PCR primer for A+B boxes mutant of human HMG1  
gcggccgc tcacttgcttttcagccttgac

FIG. 12L

SEQ ID NO:12 - forward PCR primer for B box mutant of human HMG1  
aagttcaaggatcccaatgcaaag

FIG. 12M

SEQ ID NO:13 - reverse PCR primer for B box mutant of human HMG1  
gcggccgctcaatatgcagctatatcctttc

FIG. 12N

SEQ ID NO:14 - forward PCR primer for N'+A box mutant of human HMG1  
gatgggcaaaggagatcctaag

FIG. 12O

SEQ ID NO:15 - reverse PCR primer for N'+A box mutant of human HMG1  
tcactttttgtctccctttggg

Docket/App No.: 3258.1008-001  
Title: HMGB1 Combination Therapies  
Inventors: Walter Newman }  
}

1 mgkgdpkkpr gkmssyaffv qtcreehkkk hpdasvnse fskkcserwk tmsakekgkf *rat # P07155*  
1 mgkgdpkkpr gkmssyaffv qtcreehkkk hpdasvnse fskkcserwk tmsakekgkf *mouse #AAA20508*  
1 mgkgdpkkpt gkmssyaffv qtcreehkkk hpdasvnse fskkcserwk tmsakekgkf *human #AAA64970*

A box

61 edmakadkar yeremktyip pkgetkkfk dpnapkrpps afflfcseyr pkikgehpgl *rat*  
61 edmakadkar yeremktyip pkgetkkfk dpnapkrpps afflfcseyr pkikgehpgl *mouse*  
61 edmakadkar yeremktyip pkgetkkfk dpnapkrpls afflfcseyr pkikgehpgl human

B box

121 sigdvakkig emwnntaadd kqpyekkaak lkekkekdia ayrakgpda akkgvvkaek *rat*  
121 sigdvakkig emwnntaadd kqpyekkaak lkekkekdia ayrakgpda akkgvvkaek *mouse*  
121 sigdvakkig emwnntaadd kqpyekkaak lkekkekdia ayrakgpda akkgvvkaek human

181 skkkkeeedd eedeedeeee eeeee deee dddde *rat*  
181 skkkkeeedd eedeedeeee eeeee deee dddde *mouse*  
181 skkkkeeedd eedeedeeee eeeee deee dddde *human*

FIG. 13

**FIG. 14A**

**NG\_000897 DNA (bases 150-797)**

ATGGGCAAAG GAGATCCTAA GAAGCCGACA GGCAAAATGT CATCATATGC  
ATTTTTGTG CAAACTTGTC GGGAGGAGCA TAAGAAGAAG CACCCAGATG  
CTTCAGTCAA CTTCTCAGAG TTTCTAAGA AGTGTCTAGA GAGGTGGAAG  
ACCATGTCTG CTAAAGAGAA AGGAAAATTG GAAGATATGG CAAAGGCCGA  
CAAGGCCGT TATGAAAGAG AAATGAAAAC CTATATCCCT CCCAAAGGGG  
AGACAAAAAA GAAGTTCAAG GATCCAATG CACCCAAGAG GCCTCCTTCG  
GCCTTCTTCC TCTTCTGCTC TGAGTATCGC CCAAAATCA AAGGAGAAC  
TCCTGGCCTG TCCATTGGTG ATGTTGCGAA GAAACTGGGA GAGATGTGGA  
ATAACACTGC TGCAGATGAC AAGCAGCCTT ATGAAAAGAA GGCTGCGAAG  
CTGAAGGAAA AATACGAAAA GGATATAGCT GCATATCGAG CTAAAGGAAA  
GCCTGATGCA GCAAAAAGG GAGTTGTCAA GGCTGAAAAA AGCAAGAAAA  
AGAAGGAAGA GGAGGAAGAT GAGGAAGATG AAGAGGATGA GGAGGAGGAG  
GAAGATGAAG AAGATGAAGA AGATGAAGAA GAAGATGATG ATGATGAA

**FIG. 14B**

**NG\_000897 Protein**

MGKGDPKKPT GKMSSYAFFV QTCTREEHKKK HPDASVNFSE FSKKCSERWK  
TMSAKEKGKF EDMAKADKAR YEREMKTYIP PKGETKKFK DPNAPKRLPS  
AFFLFCSEYR PKIKGEHPGL SIGDVAKKLG EMWNNTAADD KOPYEKKA  
LKEKYEKDIA AYRAKGKPDA AKKGVVKAEK SKKKKEEEE EDEEDEEEE  
EDEEDEEDE EDDDDDE

**FIG. 14C**

**AF076674 DNA (bases 1-633)**

ATGGGCAAAG GAGATCCTAA GAAGCCGAGA GGCAAAATGT CATCATATGC  
ATTTTTGTG CAAACTTGTC GGGAGGAGCA TAAGAAGAAG CACTCAGATG  
CTTCAGTCAA CTTCTCAGAG TTTCTAACAA AGTGTCTAGA GAGGTGGAAG  
ACCATGTCTG CTAAAGAGAA AGGAAAATTG GAGGATATGG CAAAGGCCGA  
CAAGACCCAT TATGAAAGAC AAATGAAAAC CTATATCCCT CCCAAAGGGG  
AGACAAAAAA GAAGTTCAAG GATCCAATG CACCCAAGAG GCCTCCTTCG  
GCCTTCTTCC TGTTCTGCTC TGAGTATCAC CCAAAATCA AAGGAGAAC  
TCCTGGCCTG TCCATTGGTG ATGTTGCGAA GAAACTGGGA GAGATGTGGA  
ATAACACTGC TGCAGATGAC AAGCAGCCTG GTGAAAAGAA GGCTGCGAAG  
CTGAAGGAAA AATACGAAAA GGATATTGCT GCATATCAAG CTAAAGGAAA  
GCCTGAGGCA GCAAAAAGG GAGTTGTCAA AGCTGAAAAA AGCAAGAAAA  
AGAAGGAAGA GGAGGAAGAT GAGGAAGATG AAGAGGATGA GGAGGAGGAA  
GATGAAGAAG ATGAAGAAGA TGATGATGAT GAA

**FIG. 14D**

**AF076674 Protein**

MGKGDPKKPR GKMSSYAFFV QTCTREEHKKK HSDASVNFSE FSNKCSERWK  
TMSAKEKGKF EDMAKADKTH YERQMKTYIP PKGETKKFK DPNAPKRPPS  
AFFLFCSEYH PKIKGEHPGL SIGDVAKKLG EMWNNTAADD KQPGEKKA  
LKEKYEKDIA AYQAKGKPEA AKKGVVKAEK SKKKKEEEE EDEEDEEEE  
DEEDEEDDDDE E

**FIG. 14E**

**AF076676 DNA (bases 1-564)**

ATGGGCAAAG GAGACCCCTAA GAAGCCGAGA GGCAAAATGT CATCATATGC  
ATTTTTGTG CAAACTTGTG GGGGAGGAGTG TAAGAAGAAG CACCCAGATG  
CTTCAGTCAA CTTCTCAGAG TTTTCTAAGA AGTGCCTCAGA GAGGTGGAAG  
GCCATGTCTG CTAAGATAA AGGAAAATTT GAAGATATGG CAAAGGTGGA  
CAAAGACCGT TATGAAAGAG AAATGAAAAC CTATATCCCT CCTAAAGGGG  
AGACAAAAAA GAAGTTCGAG GATTCCAATG CACCCAAGAG GCCTCCTTCG  
GCCTTTTGC TGTTCTGCTC TGAGTATTGC CCAAAATCA AAGGAGAGCA  
TCCTGGCCTG CCTATTAGCG ATGTTGAAA GAAACTGGTA GAGATGTGGA  
ATAACACTTT TGCAGATGAC AAGCAGCTT GTGAAAAGAA GGCTGCAAAG  
CTGAAGGAAA AATACAAAAA GGATACAGCT ACATATCGAG CTAAAGGAAA  
GCCTGATGCA GCAAAAAGG GAGTTGTCAA GGCTGAAAAA AGCAAGAAAA  
AGAAGGAAGA GGAG

**FIG. 14F**

**AF076676 Protein**

MGKGDPKKPR GKMSSYAFFV QTCREECKKK HPDASVNFSE FSKKCSEWK  
AMSAKDKGKF EDMAKVDKDR YEREMKTYIP PKGETKKKFE DSNAPKRPPS  
AFLLFCSEYC PKIKGEHPGL PISDVAKKLV EMWNNTFADD KQLCEKKA  
LKEKYKKDTA TYRAKGKPDA AKKGVVKAEK SKKKKEEE

**FIG. 14G**

**AC010149 DNA (bases 75503-76117)**

ATGGACAAAG CAGATCCTAA GAAGCTGAGA GGTGAAATGT TATCATATGC  
ATTTTTGTG CAAACTTGTG AGGAGGAGCA TAAGAAGAAG AACCCAGATG  
CTTCAGTCAA GTTCTCAGAG TTTTAAAGA AGTGCCTCAGA GACATGGAAG  
ACCATTGGT CTAAGAGAA AGGAAAATTT GAAGATATGG CAAAGGCGGA  
CAAGGCCAT TATGAAAGAG AAATGAAAAC CTATATCCCT CCTAAAGGGG  
AGAAAAAAA GAAGTTCAAG GATCCAATG CACCCAAGAG GCCTCCTTTG  
GCCTTTTCC TGTTCTGCTC TGAGTATCGC CCAAAATCA AAGGAGAAC  
TCCTGGCCTG TCCATTGATG ATGTTGTGAA GAAACTGGCA GGGATGTGGA  
ATAACACCGC TGCAGCTGAC AAGCAGTTT ATGAAAAGAA GGCTGCAAAG  
CTGAAGGAAA AATACAAAAA GGATATTGCT GCATATCGAG CTAAAGGAAA  
GCCTAATTCA GCAAAAAGA GAGTTGTCAA GGCTGAAAAA AGCAAGAAAA  
AGAAGGAAGA GGAAGAAGAT GAAGAGGATG AACAAAGAGGA GGAAAATGAA  
GAAGATGATG ATAAA

**FIG. 14H**

**AC010149 Protein**

MDKADPKKLR GEMLSYAFFV QTCQEEHKKK NPDASVKFSE FLKKCSETWK  
TIFAKEKGKF EDMAKADKAH YEREMKTYIP PKGEKKKFK DPNAPKRPL  
AFLLFCSEYR PKIKGEHPGL SIDDVVKKLA GMWNNTAAAD KQFYEKKA  
LKEKYKKDIA AYRAKGKPNS AKKRVVKAEK SKKKKEEEED EDEEQEEENE  
EDDDK

**FIG. 14I**

**AF165168 DNA (bases 729-968)**

ATGGGCAAAG GAGATCCTAA GAAGCCGAGA GGCAAAATGT CATCATGTGC  
ATTTTTGTG CAAACTGTT GGGAGGAGCA TAAGAAGCAG TACCCAGATG  
CTTCAATCAA CTTCTCAGAG TTTTCTCAGA AGTGCCAGA GACGTGGAAG  
ACCACGATTG CTAAAGAGAA AGGAAAATTT GAAGATATGC CAAAGGCAGA  
CAAGGCCAT TATGAAAGAG AAATGAAAAC CTATATAACCC

**FIG. 14J**

**AF165168 Protein**

MGKGDPKKPR GKMSSCAFFV QTCWEEHKKQ YPDASINFSE FSQKCPETWK  
TTIAKEKGKF EDMPKADKAH YEREMKTYIP

**FIG. 14K**

**XM\_063129 DNA (bases 319-558)**

AAACAGAGAG GCAAAATGCC ATCGTATGTA TTTTGTGTGC AAACCTTGTCC  
GGAGGAGCGT AAGAAGAAC ACCCAGATGC TTCAGTCAAC TTCTCAGAGT  
TTTCTAAGAA GTGCTTAGTG AGGGGGAAGA CCATGTCAGC TAAAGAGAAA  
GGACAATTG AAGCTATGGC AAGGGCAGAC AAGGCCGTT ACGAAAGAGA  
AATGAAAACA TATATCCCTC CTAAAGGGGA GACAAAAAAA

**FIG. 14L**

**XM\_063129 Protein**

KQRGKMPSYV FCVQTCPEER KKKHPDASVN FSEFSKKCLV RGKTMSAKEK  
GQFEAMARAD KARYEREMKT YIPPKGETKK

**FIG. 14M**

**XM\_066789 DNA (bases 1-258)**

ATGGGCAAAA GAGACCCTAA GCAGCCAAGA GGCAAAATGT CATCATATGC  
ATTTTTGTG CAAACTGCTC AGGAGGAGCA CAAGAAGAAA CAAACTAGATG  
CTTCAGTCAG TTTCTCAGAG TTTTCTAAGA ACTGCTCAGA GAGGTGGAAG  
ACCATGTCTG TTAAAGAGAA AGGAAAATTT GAAGACATGG CAAAGGCAGA  
CAAGGCCTGT TATGAAAGAG AAATGAAAAT ATATCCCTAC TTAAAGGGGA  
GACAAAAAA

**FIG. 14N**

**XM\_066789 Protein**

MGKRDPKQPR GKMSSYAFFV QTAQEEHKKK QLDASVSFSE FSKNCSERWK  
TMSVKEKGKF EDMAKADKAC YEREMKIYPY LKGRQK

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Title: HMGB1 Combination Therapies  
Inventors: Walter Newman

**FIG. 14O**

**AF165167 DNA (bases 456-666)**

ATGGGCAAAG GAGACCCTAA GAAGCCAAGA GAGAAAATGC CATCATATGC  
ATTTTTGTG CAAACTTGTA GGGAGGCACA TAAGAACAAA CATCCAGATG  
CTTCAGTCAA CTCCTCAGAG TTTTCTAAGA AGTGCTCAGA GAGGTGGAAG  
ACCATGCCTA CTAAACAGAA AGGAAAATTC GAAGATATGG CAAAGGCAGA  
CAGGGCCCCAT A

**FIG. 14P**

**AF165167 Protein**

MGKGDPKKPR EKMPSYAFFV QTCREAHKNK HPDASVNSSE FSKKCSERWK  
TMPTKQKGKF EDMAKADRAH